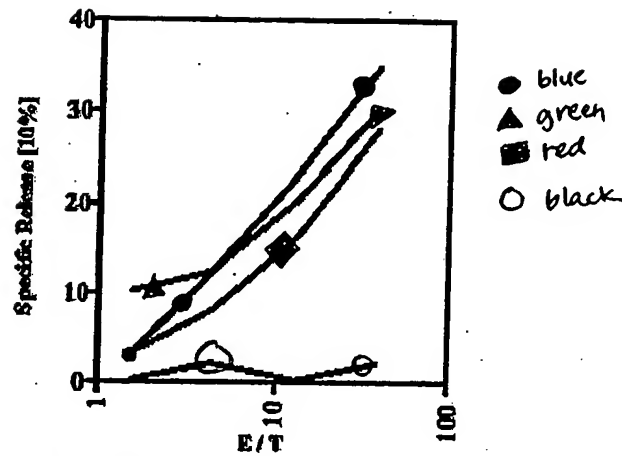


4. Balb/c spleen cells were stimulated with CS7BL/6 spleen cells. Cultures were supplemented with normal fibroblasts (blue), medium (red) or fibroblasts with CD8 (green) of mouse (A) or human (B) origin. Cultures were harvested and tested for their lytic ability towards CS7BL/6-derived target cells.

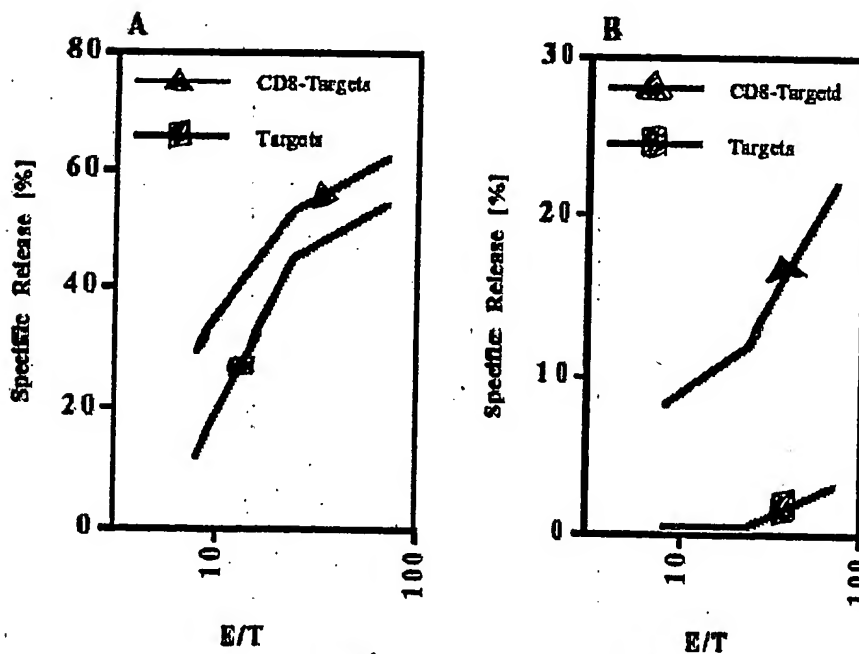
Figure 3

ANNOTATED SHEET  
SHOWING CHANGES



~~Balb/c (H-2d) mice were injected with control fibroblasts (red and green) or mCD8-transfected C57BL/6 (H-2b) derived (black and blue) fibroblasts. After two weeks animals were sacrificed, spleen cells were harvested, stimulated with C57BL/6 (H-2b) (red and black) or CBA/J (H-2k) (blue and green) spleen cells and tested for their lytic ability on EL4 (H-2b) (red and black) or S.AKR (H-2k) (blue and green) target cells.~~

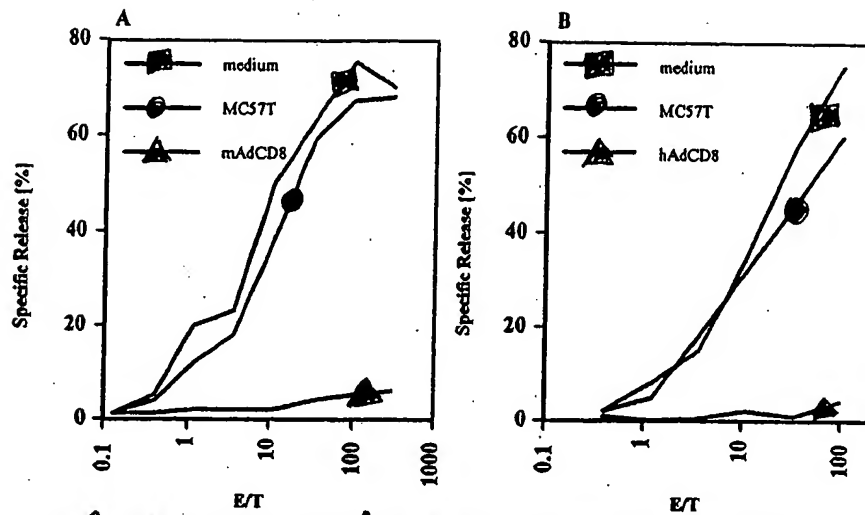
Figure 4



~~Target cells (green) or CD8-expressing targets (red) were tested for their susceptibility to lysis by alloreactive T cells (A) or by antigen-specific CTLs (B).~~

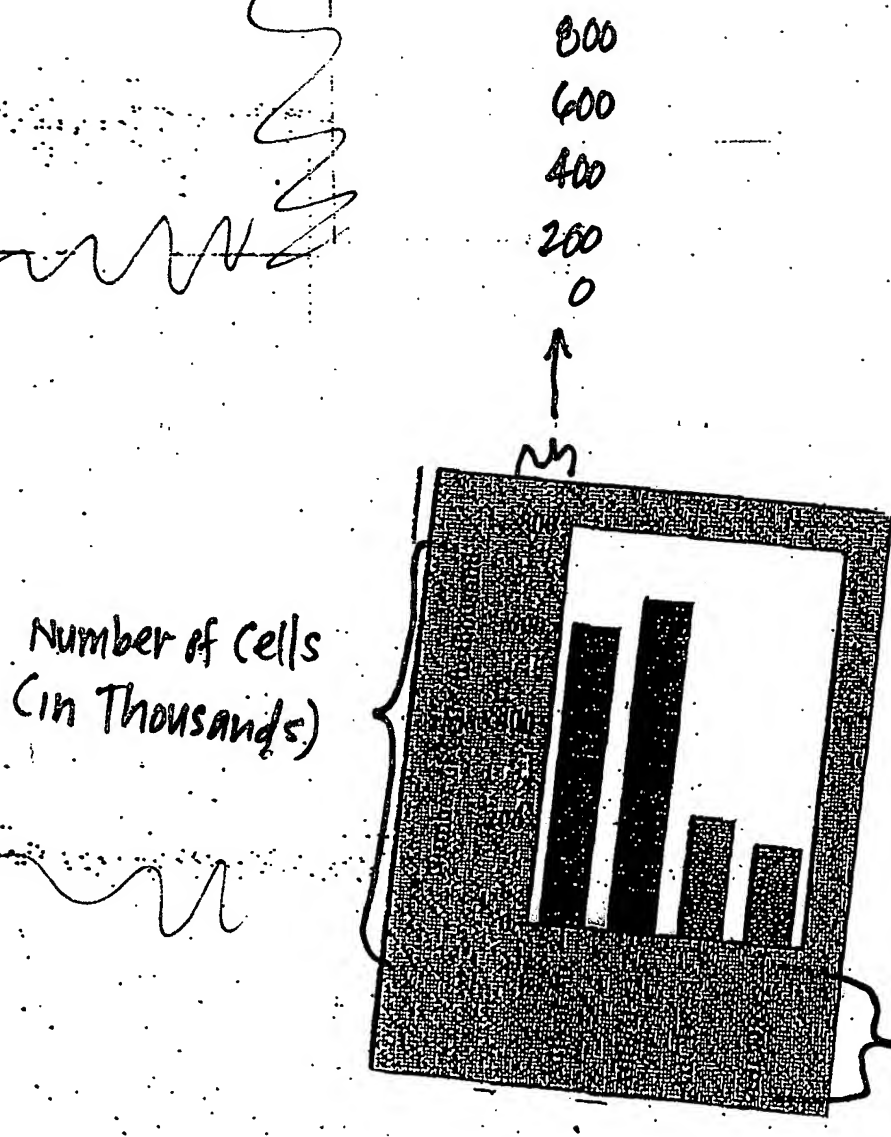
Figure 5

# ANNOTATED SHEET SHOWING CHANGES



~~circle~~ MLCs (Balb/c anti-C57BL/6) were set up in the presence of normal fibroblasts (blue) and fibroblasts transduced with mAdCD8 (A, green) or hAdCD8 (B, green). No fibroblasts were added to control cultures (red). The lytic activity of these cultures towards an O57BL/6-derived target was determined at the end of the culture period.

Figure 6



Adβgal #1  
Adβgal #2  
AdCDB #1  
AdCDB #2

FIGURE 7

ANNOTATED SHEET  
SHOWING CHANGES

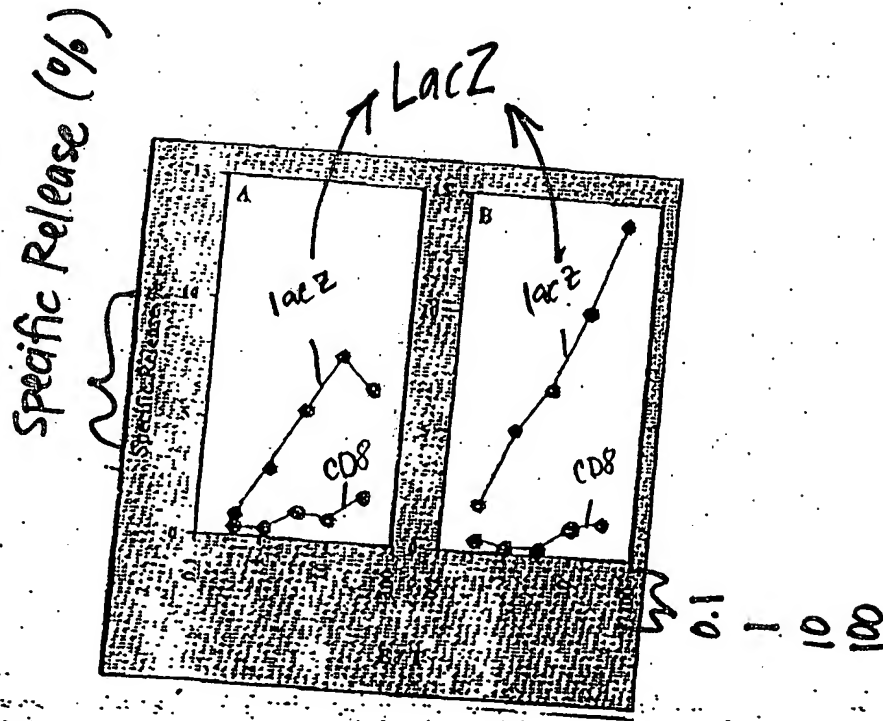
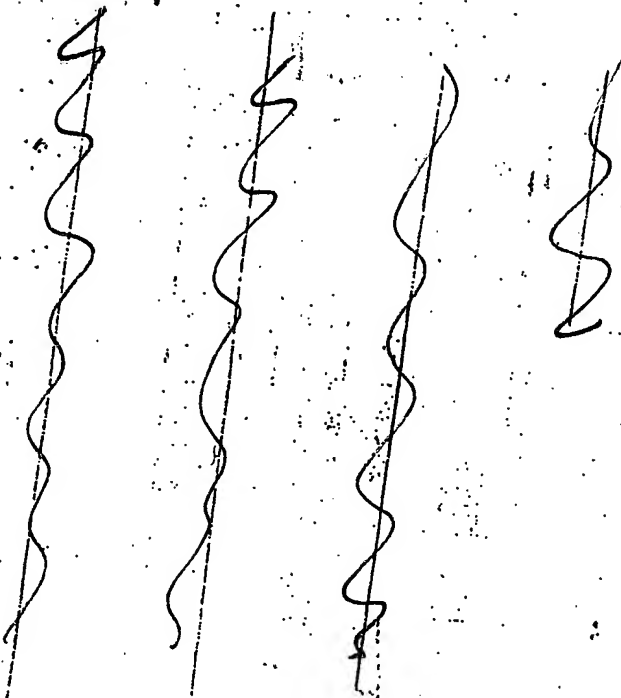
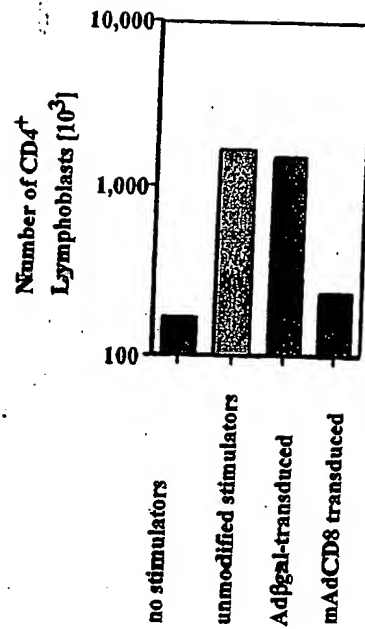


FIGURE 8



ANNOTATED SHEET  
SHOWING CHANGES



~~3x10<sup>6</sup> C7BI/6 spleen cells were incubated with 1x10<sup>6</sup> (or no) stimulator cells, transduced as indicated. After 4 days the cultures were analyzed for presence CD4<sup>+</sup> T lymphoblasts by immunofluorescence.~~

Figure 9

ANNOTATED SHEET  
SHOWING CHANGES

FIGURE 10A

~~Infected Cells: MC57T Fibroblasts~~  
~~Panel 1: Mook-Infection; Panel 2: Infection with hAdCD8~~

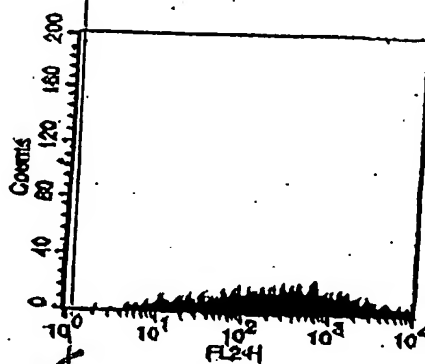
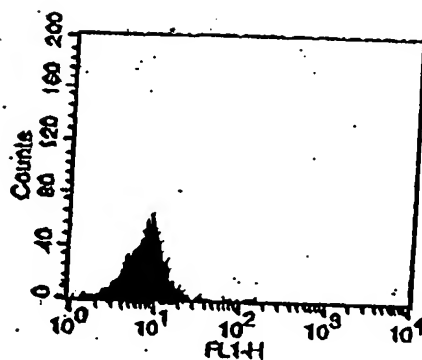
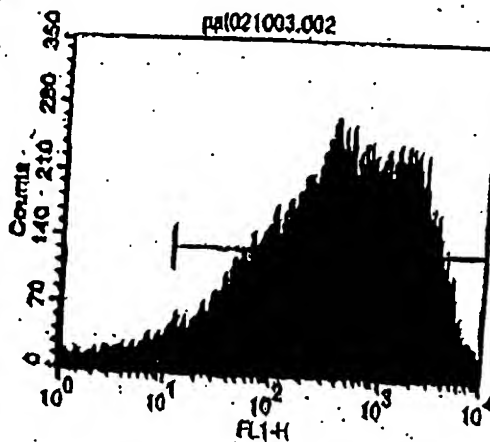
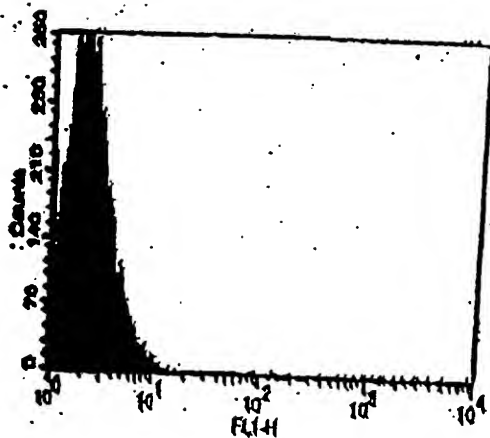


FIGURE 10B

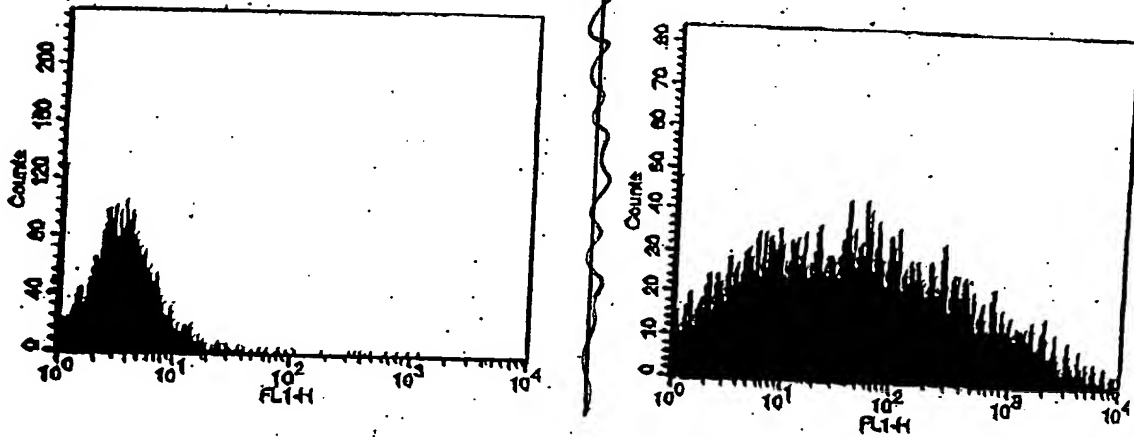
~~Infected Cells: MC57T Fibroblasts~~  
~~Panel 1: Mook-Infection; Panel 2: Infection with mAdCD8~~



ANNOTATED SHEET  
SHOWING CHANGES

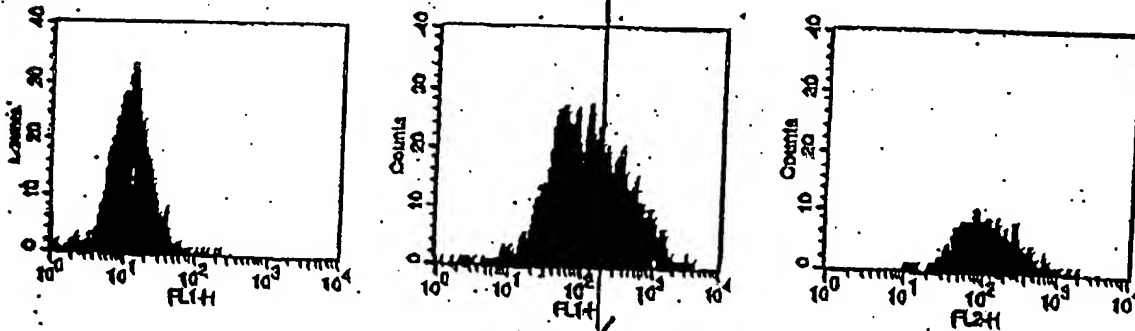
**FIGURE 10C**

~~Infected Cells: Balb/c unselected bone marrow cells;  
Panel 1: Infection with lacZ Adenoviral Vector (AdLacZ);  
Panel 2: Infection with mAdCD8~~



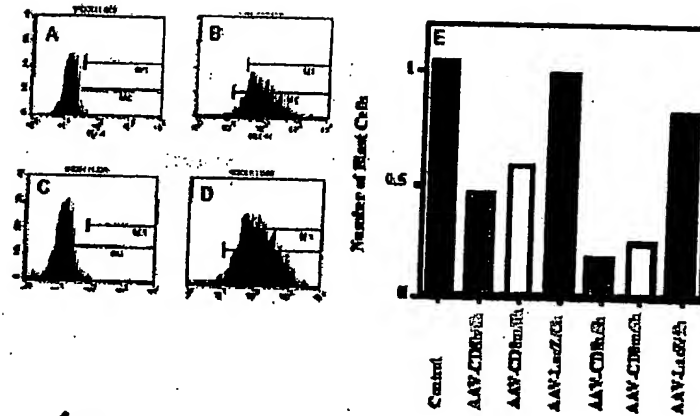
**FIGURE 10D**

~~Infected Cells: MC57T Fibroblasts  
Panel 1: Mock-Infection;  
Panel 2: Infection with pAAV-mCD8;  
Panel 3: Infection with pAAV-hCD8~~





ANNOTATED SHEET  
SHOWING CHANGES

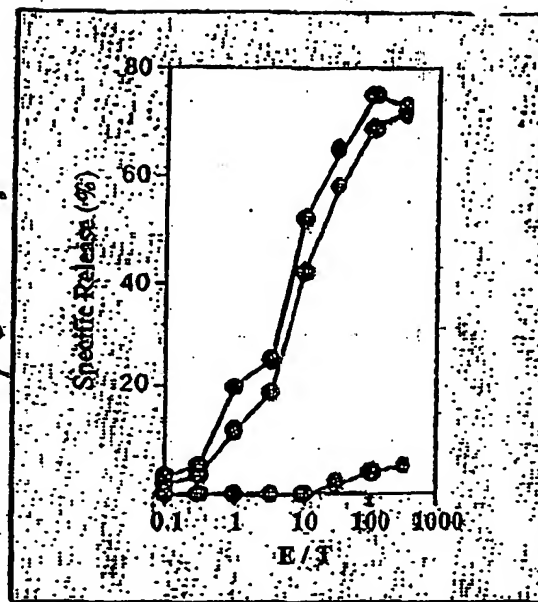


Fibroblasts were transduced with pAAVCD8 (B) or hAAVCD8 (D) or mock-infected (A and C). Surface expression of CD8 was detected by surface immunofluorescence (A through D). MLCs (Balb/c anti-C57BL/6) were set up in the presence of these fibroblasts that had been cultured for 0 or 5 hours after transduction before they were added to the MLCs. At end of cultures, the number of lymphoblasts was determined on a fluorescence activated cell analyzer.

Figure 11

ANNOTATED SHEET  
SHOWING CHANGES

Specific Release (%)

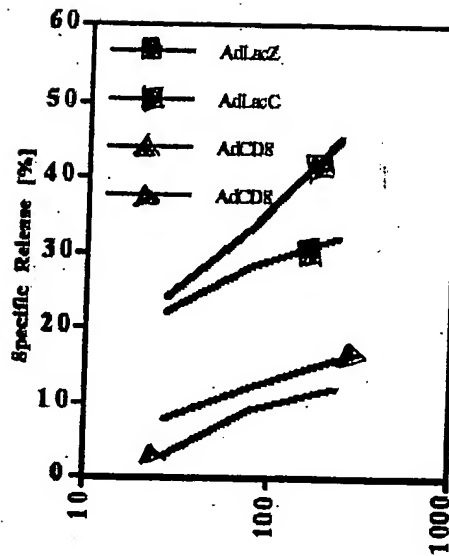


0.1 1 10 100 1000

E/T

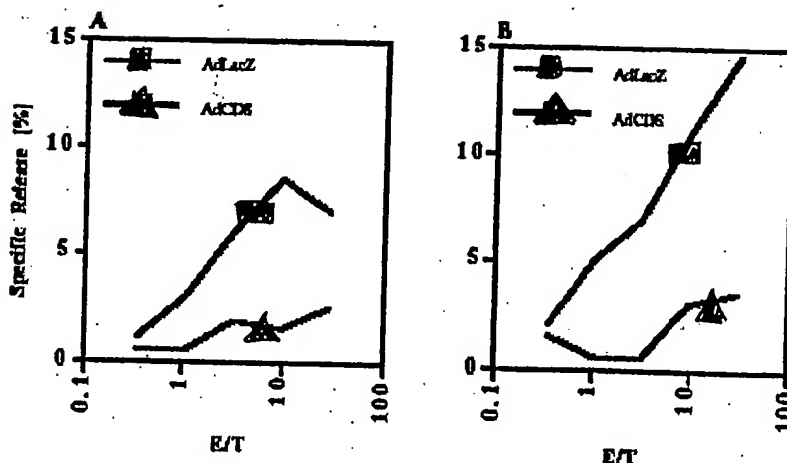
FIGURE 12

ANNOTATED SHEET  
SHOWING CHANGES



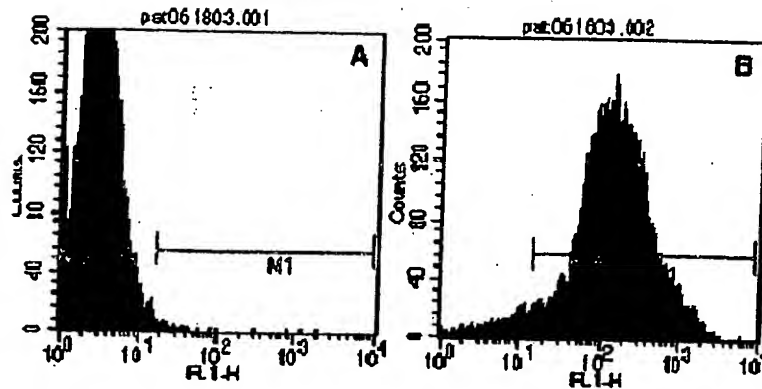
~~triangle~~ <sup>E/T</sup> Balb/c mice were immunized with AdLacZ (green) or mAdCD8 (red). Their spleen cells were cultured in the presence of AdLacZ and tested for specific lytic activity against AdLacZ-infected syngeneic P815 target cells.

Figure 13



~~triangle~~ <sup>E/T</sup> (A) C57BL/6 animals were immunized with AdLacZ (red) or mAdCD8 (green). Their lytic activity of their spleen cells towards syngeneic AdLacZ BL4 target cells was tested. (B) Such animals were re-immunized with AdLacZ prior to testing their lytic activity against AdLacZ-infected BL4 targets.

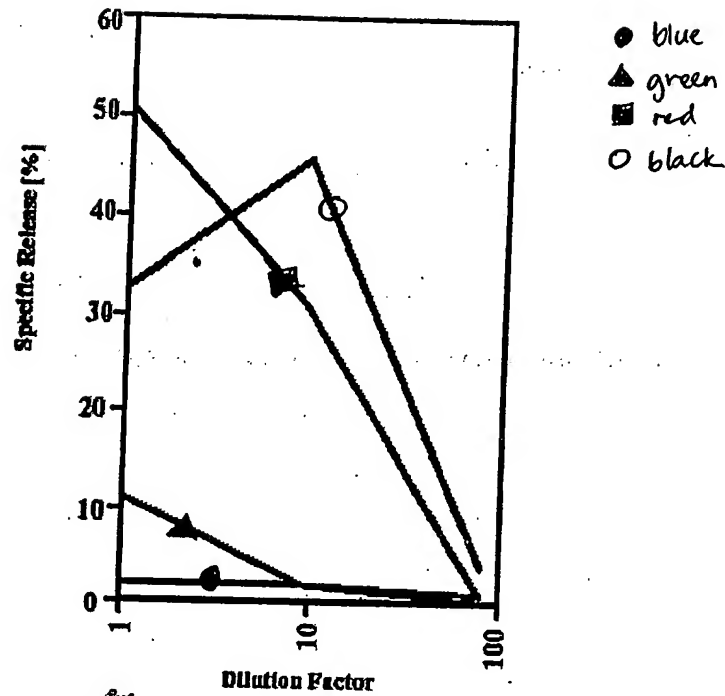
ANNOTATED SHEET  
SHOWING CHANGES



~~Single cell suspensions were prepared from newborn hearts. The heart muscle cells were transduced with mAdCD8 (B) or mock-infected, cultured for 48 hours and stained for the surface expression of CD8.~~

Figure 15

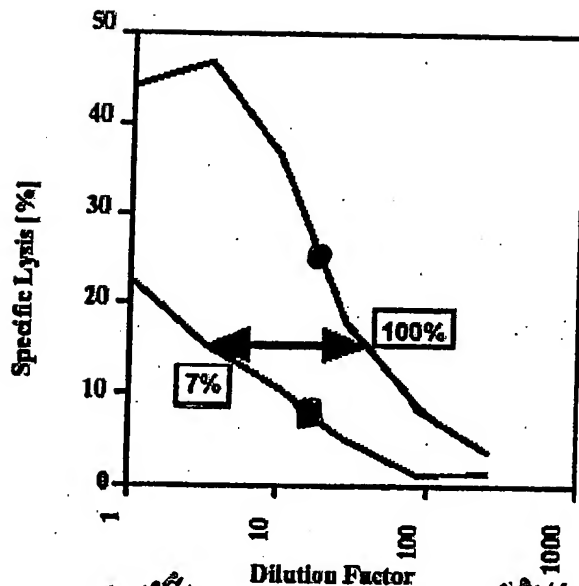
ANNOTATED SHEET  
SHOWING CHANGES



*Square*  
Newborn C57BL/6 hearts were infected  
with  $10^8$  (red),  $5 \times 10^7$  (green),  $10^7$  (blue) PFU AdCD8  
or mock-infected (black). Thirty-five days after  
transplantation into BAL B/c recipients, the activity of  
the lytic activity of activated recipient T cells was  
tested on donor-type target cells.  
*per chart*

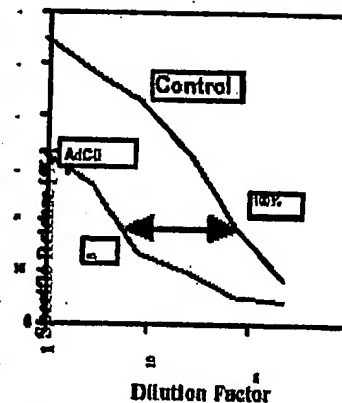
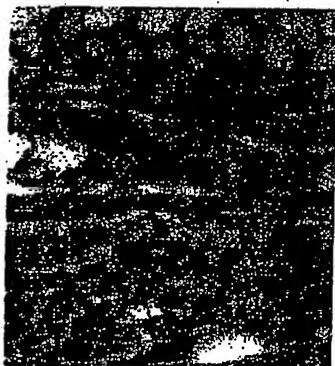
Figure 16

ANNOTATED SHEET  
SHOWING CHANGES



Newborn C57BL/6 hearts were infected with AdCD8 (red) or mock-infected (black). Thirty-eight days after transplantation into BALB/c recipients, the activity of the lytic activity of activated recipient T cells was tested on donor-type target cells.

Figure 17



Animal: #725

C57BL/6 hearts infected with rAdCD8 (treated) or mock-infected (control) were transplanted into Balb/c mice. After 52 days, the animals were sacrificed and the tissue was stained (HE) and the lytic activity of recipient T cells was tested on donor-type target cells.

Figure 18

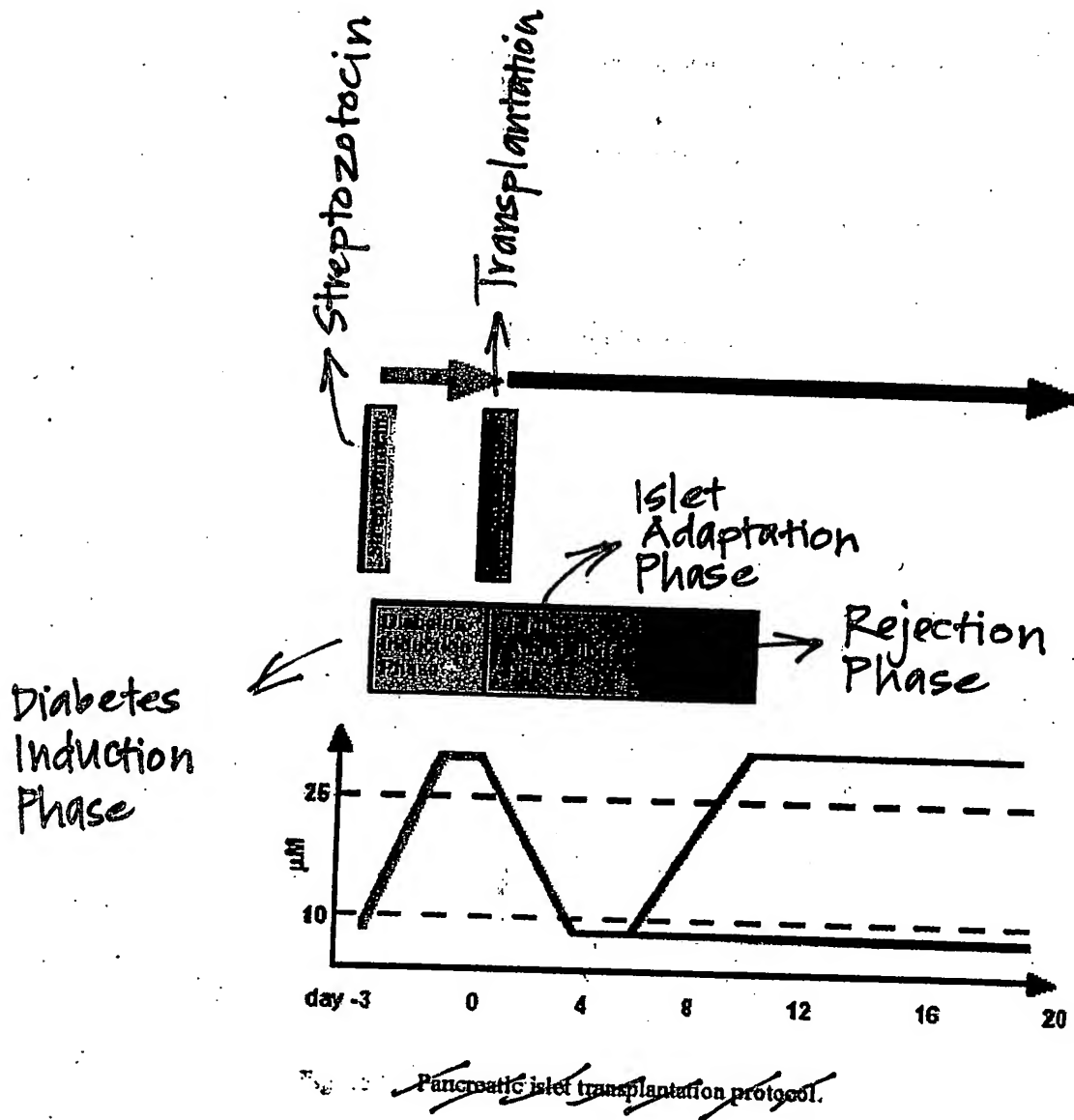
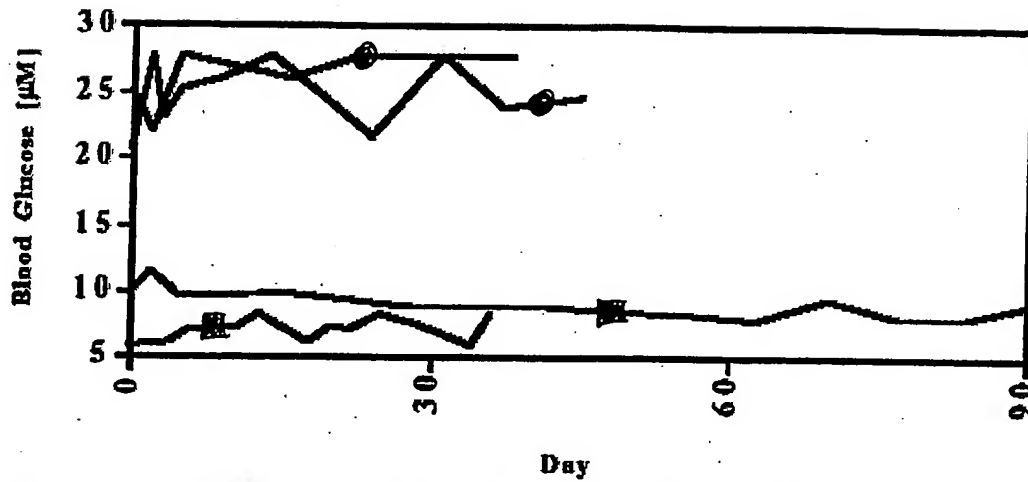


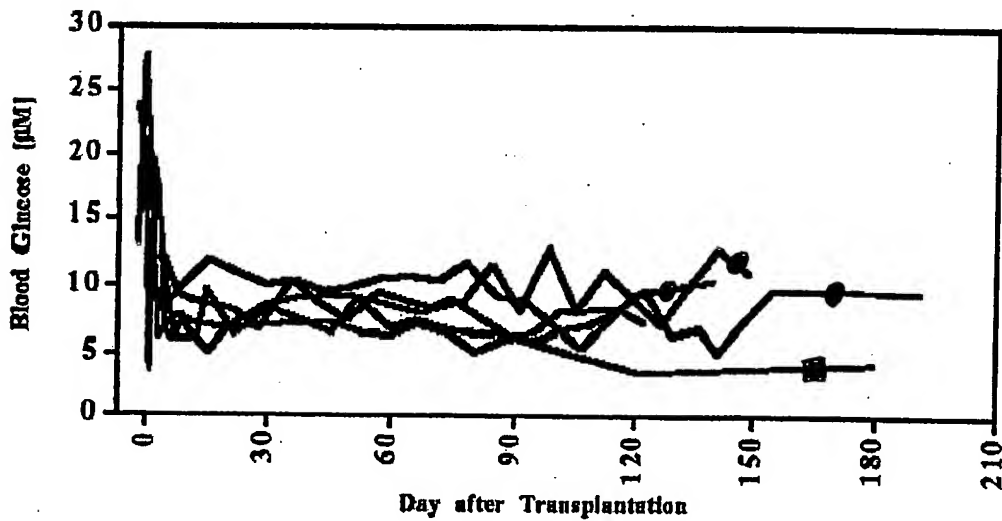
Figure 19

ANNOTATED SHEET  
SHOWING CHANGES



Blood glucose levels in normal (red) and Streptozotocin-treated (blue) mice.  
*Square* *Circle*

Figure 20

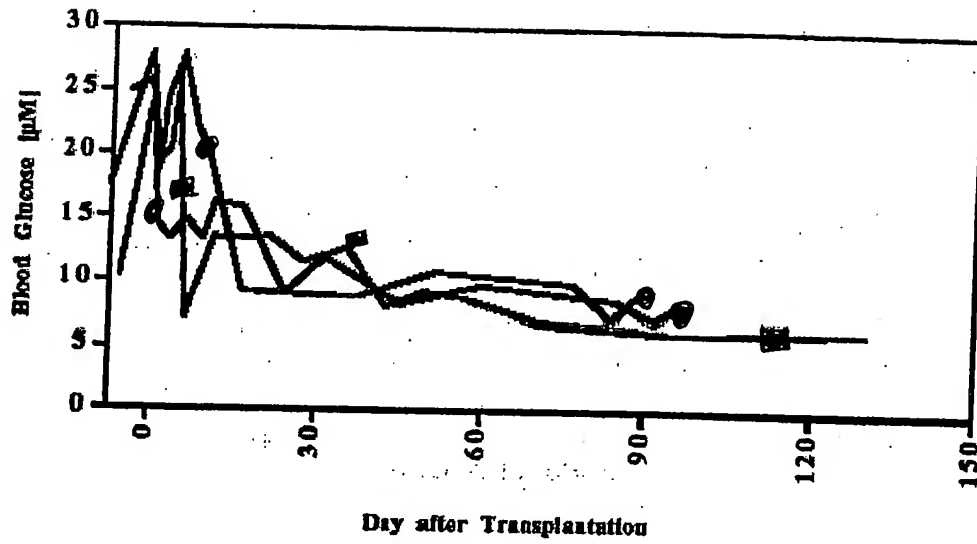


Syngeneic pancreatic islet transplants performed in Balb/c (red) and in C57BL/6 (blue) mice.  
*Square* *Circle*

Figure 21



ANNOTATED SHEET  
SHOWING CHANGES



Transplantation of syngeneic mAdCD8-transduced pancreatic islets  
harvested from Balb/c (blue) or C57BL/6 (red) mice.  
circle square

Figure 22

ANNOTATED SHEET  
SHOWING CHANGES

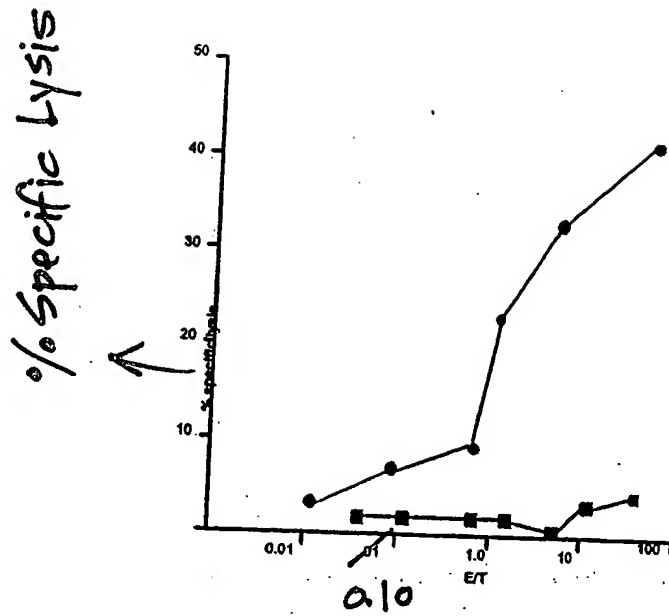
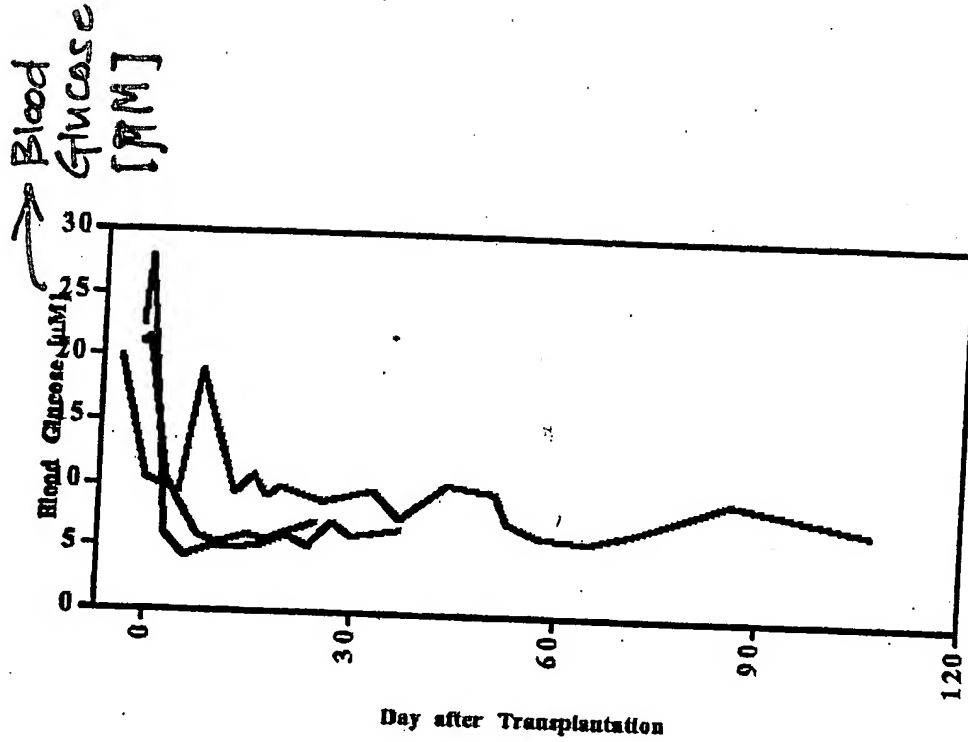


Figure 24

ANNOTATED SHEET  
SHOWING CHANGES



mAdCD8-transduced C57BL/6 pancreatic islets were transplanted into Balb/c recipient mice.

Figure 25

**This Page is Inserted by IFW Indexing and Scanning  
Operations and is not part of the Official Record**

## **BEST AVAILABLE IMAGES**

Defective images within this document are accurate representations of the original documents submitted by the applicant.

Defects in the images include but are not limited to the items checked:

- ☐ **BLACK BORDERS**
- ☐ **IMAGE CUT OFF AT TOP, BOTTOM OR SIDES**
- ☐ **FADED TEXT OR DRAWING**
- ☐ **BLURRED OR ILLEGIBLE TEXT OR DRAWING**
- ☐ **SKEWED/SLANTED IMAGES**
- ☐ **COLOR OR BLACK AND WHITE PHOTOGRAPHS**
- ☐ **GRAY SCALE DOCUMENTS**
- ☒ **LINES OR MARKS ON ORIGINAL DOCUMENT**
- ☐ **REFERENCE(S) OR EXHIBIT(S) SUBMITTED ARE POOR QUALITY**
- ☐ **OTHER:** \_\_\_\_\_

**IMAGES ARE BEST AVAILABLE COPY.**

**As rescanning these documents will not correct the image problems checked, please do not report these problems to the IFW Image Problem Mailbox.**